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"Intraosseous Erythropoietin for Acute Tissue Protection in Battlefield Casualties Suffering Hypovolemic Shock"
PRINCIPAL INVESTIGATOR:
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#### 13. SUPPLEMENTARY NOTES

#### 14. ABSTRACT

The project compared in a swine model of hemorrhagic shock (HS) the effects of vasopressin (VP) infusion and low-volume fluid resuscitation using normal saline (NS) on 72 hour survival, with erythropoietin (EPO) administration and HS severity as confounders. Twenty-four male pigs (36-41 kg) were anesthetized with isoflurane and nitrous oxide in oxygen and instrumented to assess hemodynamic function and advance a 23-F cannula into the right atrium for blood withdrawal (BW) and blood reinfusion using a custom developed system that allowed modeling spontaneous bleeding as a mono-exponential decay function. Twenty-four pigs were randomized 2:1 to receive an intraosseous infusion of VP (0.04 U/kg·min<sup>-1</sup>) or vehicle control starting 7 minutes into BW until the start of blood reinfusion at minute 210. Pigs were also randomized 1:1 to receive NS (half the amount of BW) or no fluids. Pigs assigned to VP were also randomized 1:1 to EPO (1,200 U/kg) or vehicle control and to have 65% or 75% of their blood volume withdrawn. Randomization proceeded by blocks ensuring balanced distribution in all the subsets. Survival analysis to 72 hours showed that survival was influenced by VP and NS but not by EPO or HS severity with the highest survival rate in the group VP-NS (100%) followed by VP-noNS (37.5%), noVP-NS (25%), and noVP-noNS (0%) with a high overall statistical significance (*p*=0.009 by the log-rank test) with each subset different than VP-NS by the Holm-Sidak's test. Accordingly, these findings support an approach to the initial management of severe HS in the field that entails early initiation of VP infusion through the intraosseous route followed by low-volume fluid resuscitation (e.g., small boluses of normal saline).

# 15. SUBJECT TERMS

Shock, Hemorrhagic; Resuscitation; Erythropoietin; Vasopressin; Swine; Polytrauma and Blast Injury; Maintenance of Hemodynamic Stability; Low-Volume Fluid Resuscitation

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#### INTRODUCTION

Hemorrhagic shock after penetrating trauma in the battlefield accounts for a high percentage of potentially survivable injuries. A report on 4,596 battlefield fatalities between October 2001 and June 2011 in Operation Iraqi Freedom and Operation Enduring Freedom showed that 87% of such fatalities occurred before arrival to a medical treatment facility with 24% deemed potentially survivable. Of these potentially survivable injuries, 91% were associated with hemorrhage. Control of

#### **Abbreviations:**

BR = Blood reinfusion

EPO = Erythropoiet in

VP = Vasopressin

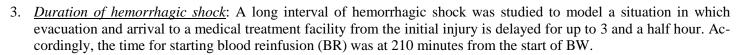
NS = Normal saline BW = Blood withdrawal

bleeding in compressible areas is highly effective and can save lives. However, severe hemorrhage can compromise vital organ perfusion and survivability. Fluid resuscitation is hemodynamically effective but is logistically constrained in the battlefield and not free of complications. In large quantities, fluids can precipitate coagulopathy by dilution and hypothermia and dislodge freshly formed clots. Similar considerations apply to hemorrhage associated with trauma in civilian populations. We have previously shown in a swine model of hemorrhagic shock – while investigating the effects of erythropoietin (EPO) – that vasopressin (VP) infusion promotes hemodynamic stability. During the reported period, we investigated whether VP infusion with or without low-volume fluid resuscitation using normal saline (NS) could improve survival from severe and protracted hemorrhagic shock produced by removing 65% or 75% of the animal's blood volume. Consistent with the initial hypothesis of this project, i.e., that EPO could ameliorate tissue injury; we also included administration of EPO in this series.

#### **BODY**

The experiments were designed based on findings reported last year (after having completed three series of experiments) refocusing the work on the potential role of VP and low-volume fluid resuscitation for hemodynamic stabilization under conditions of severe hemorrhagic shock maintaining the emphasis on battlefield relevance and constraints for management in far-forward scenarios. Accordingly, we modified our hemorrhagic shock protocol as described below using the swine preparation depicted in Figure 1:

- 1. <u>Simulated bleeding pattern</u>: The pattern of acute blood loss was changed from a linear function to a monoexponential decay function to more closely model bleeding after traumatic injury (i.e., high initial bleeding with progressively slower rates as blood pressure drops and hemostatic mechanisms are endogenously activated and/or externally applied). The approach involved additional software development by us that enabled entering any mathematical function of our choosing into our closed-loop system to drive the flow removal pattern with a precision of < 1 ml/min.
- 2. <u>Blood volume removed</u>: Two target levels of blood withdrawal (BW) were studied; 65% and 75% of the estimated blood volume. It is pertinent to highlight that in
  - our previous hemorrhagic shock series the highest amount of BW was 65% of the blood volume. Thus, for the first time experiments were conducted withdrawing 75% of the estimated blood volume. The same mono-exponential decay function was used for both BW targets, stopping after 60 minutes for the BW 65% and after 80 minutes for BW 75%



Pursue of findings originated from the previous series prompted the change in the protocol as described below:

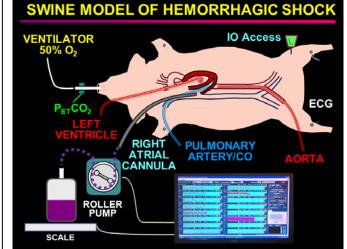
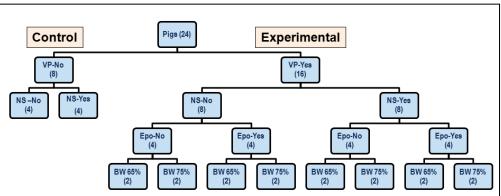


Figure 1: Rendition of our swine model of hemorrhagic shock depicting the intraosseous access for drug delivery into the left tibia; vascular catheterization to monitor left ventricular pressure, aortic pressure, pulmonary artery pressure, right atrial pressure, and cardiac output; right atrial cannulation for blood withdrawal and blood reinfusion programmatically controlled by custom developed software guided by gravimetric measurement of blood volume; and mechanical ventilation with measurement of the expired end-tidal PCO<sub>2</sub>.

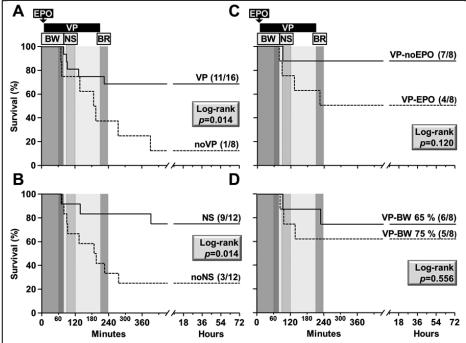
<u>Vasopressin effect</u>: As reported previously, the addition of vasopressin infusion in a previous series dramatically improved initial resuscitation and subsequent 72-hour survival relative to a previous series in which only fluid resuscitation with normal saline (NS) was used for stabilization before BR. The effect of vasopressin on these outcomes was associated with higher cardiac index, prompting us to speculate that VP could act in part by enhancing venous return. Because these observations were made in separate series, the question regarding the efficacy and hemodynamic effects of VP could not be directly addressed. Accordingly, for this series animals were randomized 2:1 to VP or noVP. VP was infused intraosseously at a constant rate of 0.04 U/kg·min<sup>-1</sup> from minute 7 of BW to minute 210; i.e., the start of BR.

<u>Fluid resuscitation</u>: The preceding series also showed that in the presence of vasopressin, initial resuscitability and 72-hour survival could be achieved successfully administering a reduced amount of NS corresponding to half the amount of BW or no fluids at all. Thus, for the present series we examined whether NS could be required under the more severe hemorrhagic shock conditions modeled. The approach involved starting the series without fluid resuscitation and performing an interim analysis halfway through the series (i.e., after completion of 12 experiments) to assess the needs for fluid resuscitation. The interim analysis showed a 72-hour survival of only 3 of 12 animals (25%) prompting the inclusion in the second half of the series the administration of NS from minute 90 to minute 120 in amount corresponding to half the volume of BW. The relatively small amount given was effective resulting in 9 out 12 animals (75%) surviving the 72 hour post-resuscitation interval.

Erythropoietin (EPO): In keeping with the original objective of our grant but incorporating knowledge gained from the



**Figure 2**: Experimental groups and subsets. VP, vasopressin; NS, normal saline; Epo, erythropoietin; BW, blood withdrawn.



**Figure 3**: Kaplan- Maier survival curves analyzed using the log-rank test to assess the independent effects of normal saline (NS), vasopressin (VP), erythropoietin (EPO), and HS severity BW, blood withdrawn (BW); BR, blood reinfusion.

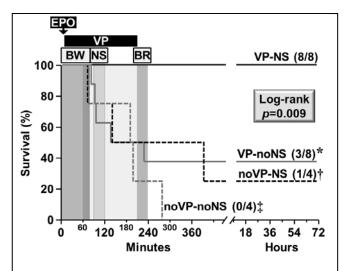
previous series, in which EPO had no effect on resuscitation and 72-hour survival but attenuated aortic lactate increases and organ dysfunction under conditions of severe but survivable hemorrhagic shock, we randomized in the subset of animals receiving VP to also receive EPO or vehicle control given – like in the preceding series – as a single intraosseous bolus dose of 1,200 U/kg after 10% of the blood volume had been removed (6 minutes from the start of BW).

<u>Hemorrhagic shock severity</u>: The effect of increasing the hemorrhagic shock severity was investigated only in the subset of animals receiving vasopressin by randomizing the animals to a severity of BW 65% or BW 75%.

Figure 2 summarizes the various groups and subsets investigated.

### **Survival Effect**

Separate Kaplan-Maier survival analyses using the log-rank test were performed for each of the main factors while maintaining a balanced distribution of the remaining factors. As shown in Figure 3A, vasopressin was associated with statistically significant higher survival (68.8% vs 12.5%, p=0.014). The favorable effect occurred even though half of the animals in the vasopressin group, but none in controls, were subjected to more severe hemorrhagic shock BW (75% of blood volume). Administration of NS was also associated



**Figure 4**: Kaplan- Maier survival curves comparing effects of vasopressin with normal saline (VP-NS), VP without NS (VP-noVP), no VP with NS (noVP-NS), and neither VP nor NS (noVP-noNS). The overall statistical significance was assessed by the log-rank test with all pairwise comparisons assessed by the Holm-Sidak's test demonstrating significantly lower survival for VP-noNS (\*p = 0.035), noVP-NS (†p = 0.0198), and noVP-noNS (‡p = 0.001) compared with VP-NS. EPO, erythropoietin; BW, blood withdrawn; BR, blood reinfusion.

with higher 72-hour survival (25% vs 75%, p=0.014) as shown in Figure 3B. The use of EPO (Figure 3C) was associated with lower survival; however, the difference was not statistically significant. The increase in hemorrhagic shock severity had only a minimal and statistically insignificant effect on survival (Figure 3D).

Analysis of the combined effects of VP and NS on 72 hour survival is shown in Figure 4 demonstrating the highest survival rate associated with the combination of VP and NS and the worse with the absence of VP and NS.

A multiple linear regression analysis assessing the combined effects of vasopressin (yes[1]/No[0]), NS (Yes[1]/No[0]), erythropoietin (Yes[1]/No[0]), and hemorrhagic shock severity (65%[0]/75%[0]) on duration of survival (hours) demonstrated a statistically significant (p<0.001) predictive effect of these independent variables with an R<sup>2</sup> = 0.632:

SURVIVAL (HOURS) = -5.7 + (55.9 X VASOPRESSIN) + (35.0 X NS) - (26.1 X ERYTHROPOIETIN) - (9.0 X SEVERITY)

However, only vasopressin (p<0.001), NS (p=0.002), and erythropoietin (p=0.040), but not severity (p=0.455) contributed statistically to the model.

# **Hemodynamic and Metabolic Effects**

The factorial design of the study allowed separate analyses of the effects of NS and the effects of NS on hemodynamic and metabolic variables as shown in Figures 5 through Figure 8. VP as expected increased systemic vascular resistance through the interval of blood withdrawal and that this effect was associated with a reduction in cardiac index and also a reduction in heart rate (Figure 5). These hemodynamic effects were associated with a lower end-tidal PCO<sub>2</sub> (P<sub>ET</sub>CO<sub>2</sub>) and more intense lactic acidosis attributed to supply-dependent reduction in oxygen consumption (Figure 6). The effect, however, did not compromise survival. We interpreted these effects as indicative of a survival benefit prompted by redistribution of blood flow toward the heart by VP with likely reduction in oxygen requirements associated with a lower heart rate know to be an effect of VP modulating the baroreflex response. Low-volume fluid resuscitation was associated with an increase in left ventricular stroke work index and cardiac index – likely an effect on preload – that helped increase the aortic pressure and reduce the systemic vascular resistance (Figure 7). Metabolically, low-volume fluid resuscitation helped reduce lactic acidosis (Figure 8).

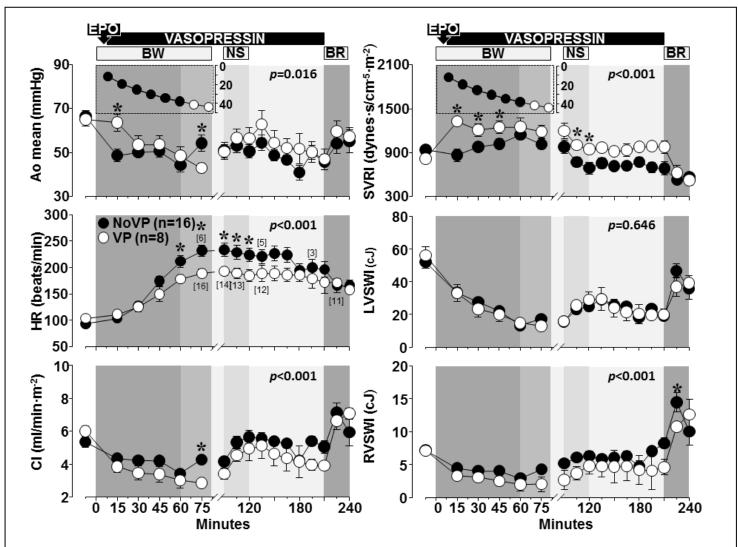
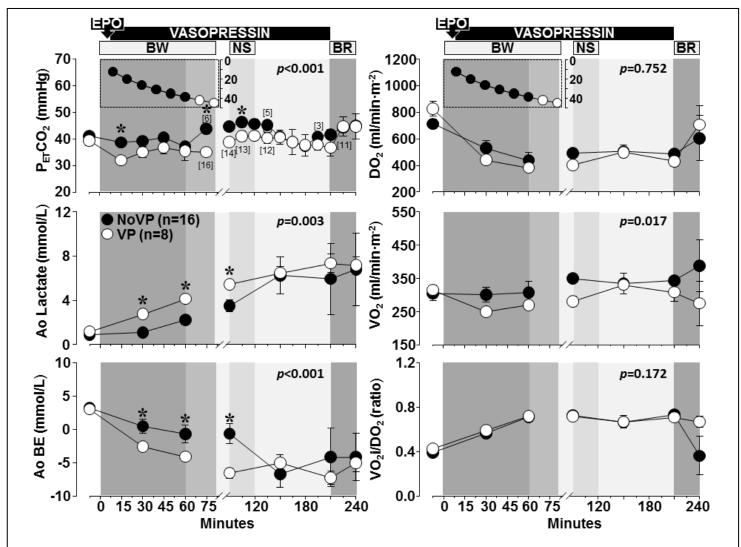


Figure 5: Effects of VP (open circles, n = 16) compared with vehicle control (closed circles, n = 8) on hemodynamic and myocardial function. Numbers in brackets indicate when the number of animals decreased from the preceding time point consequent to death. EPO, erythropoietin; VP, vasopressin; BW, blood withdrawal; HS, hemorrhagic shock; NS, normal saline; BR, blood reinfusion. The inset depicts the time course of the blood withdrawal (ml/kg) ending at 60 minutes in the 65% BW subset and at 80 minutes in the 75% BW group. Ao, aortic pressure; HR, heart rate; CI, cardiac index; SVRI, systemic vascular resistance index; LVSWI, left ventricular stroke work index; RVWI, right ventricular stroke work index. Values are shown as mean  $\pm$  SEM. Differences between groups were analyzed by two-way repeated measures ANOVA. There was an overall statistically significant treatment effect for Ao mean, (p = 0.016); heart rate, (p < 0.001); CI, (p < 0.001); SVRI (p < 0.001) and RVWI (p < 0.001). There was not statistically significant interaction between treatment and time. \* $p \le 0.05$ , denote statistically significant differences between groups at the specified time points using the Holm-Sidak test for multiple comparisons.



**Figure 6**: Effects of VP (open circles, n=16) compared with vehicle control (closed circles, n=8) on  $P_{ET}CO_2$ , lactic acidosis, and systemic oxygen metabolism. Numbers in brackets indicate when the number of animals decreased from the preceding time point consequent to death. EPO, erythropoietin; VP, vasopressin; BW, blood withdrawal; HS, hemorrhagic shock; NS, normal saline; BR, blood reinfusion. The inset depicts the time course of the blood withdrawal (ml/kg) ending at 60 minutes in the 65% BW subset and at 80 minutes in the 75% BW group.  $P_{ET}CO_2$ , end-tidal carbon dioxide; Ao, aortic; BE, base excess; DO<sub>2</sub>, oxygen delivery index; VO<sub>2</sub>, oxygen consumption index; VO<sub>2</sub>/DO<sub>2</sub>, oxygen consumption and delivery ratio. Values are shown as mean  $\pm$  SEM. Differences between groups were analyzed by two-way repeated measures ANOVA. There was an overall statistically significant treatment effect for  $P_{ET}CO_2$ , (p < 0.001); Ao lactate, (p = 0.003); Ao BE, (p < 0.001) and VO<sub>2</sub> (p = 0.017). There was an overall statistically significant interaction between treatment and time for Ao BE (p = 0.040). \* $p \le 0.05$ , denote statistically significant differences between groups at the specified time points using the Holm-Sidak test for multiple comparisons.

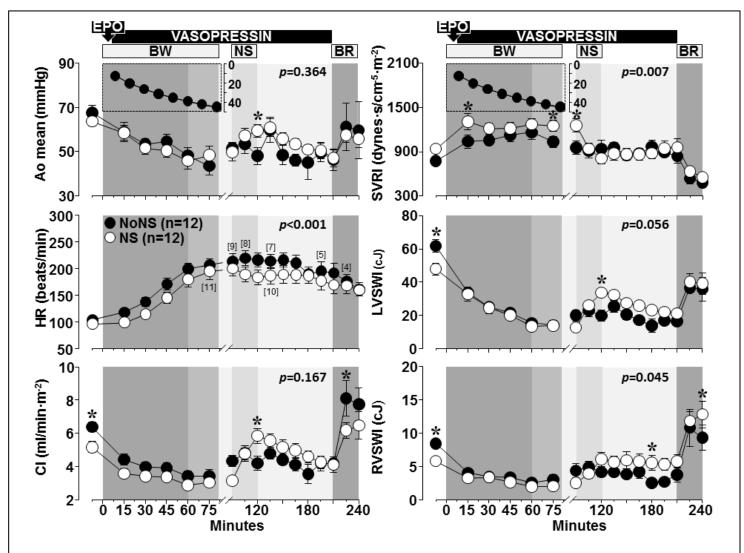
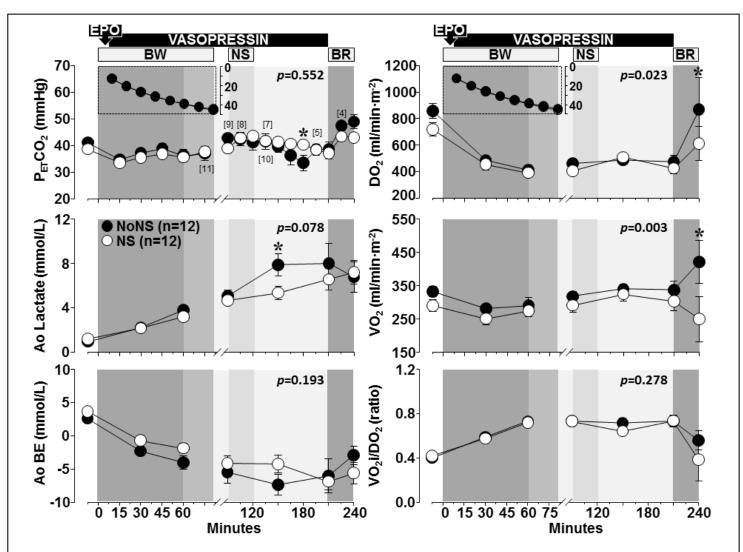


Figure 7: Effects of NS (open circles, n = 12) compared with vehicle control (closed circles, n = 12) on hemodynamic and myocardial function. Numbers in brackets indicate when the number of animals decreased from the preceding time point consequent to death. EPO, erythropoietin; BW, blood withdrawal; HS, hemorrhagic shock; NS, normal saline; BR, blood reinfusion. The inset depicts the time course of the blood withdrawal (ml/kg) ending at 60 minutes in the 65% BW subset and at 80 minutes in the 75% BW group. Ao, aortic pressure; HR, heart rate; CI, cardiac index; SVRI, systemic vascular resistance index; LVSWI, left ventricular stroke work index; RVWI, right ventricular stroke work index. Values are shown as mean  $\pm$  SEM. Differences between groups were analyzed by two-way repeated measures ANOVA. There was an overall statistically significant treatment effect for heart rate, (p < 0.001); SVRI (p = 0.007) and RVWI (p = 0.045). There was an overall statistically significant interaction between treatment and time for CI (p < 0.001), LVSWI (p = 0.007) and RVWI (p = 0.009). \* $p \le 0.05$ , denote statistically significant differences between groups at the specified time points using the Holm-Sidak test for multiple comparisons.



**Figure 8**: Effects of NS (open circles, n = 12) compared with vehicle control (closed circles, n = 12) on  $P_{ET}CO_2$ , lactic acidosis, and systemic oxygen metabolism. Numbers in brackets indicate when the number of animals decreased from the preceding time point consequent to death. EPO, erythropoietin; BW, blood withdrawal; HS, hemorrhagic shock; NS, normal saline; BR, blood reinfusion. The inset depicts the time course of the blood withdrawal (ml/kg) ending at 60 minutes in the 65% BW subset and at 80 minutes in the 75% BW group.  $P_{ET}CO_2$ , end-tidal carbon dioxide; Ao, aortic; BE, base excess;  $DO_2$ , oxygen delivery index;  $VO_2$ , oxygen consumption index;  $VO_2/DO_2$ , oxygen consumption and delivery ratio. Values are shown as mean  $\pm$  SEM. Differences between groups were analyzed by two-way repeated measures ANOVA. There was an overall statistically significant treatment effect for  $DO_2$  (p = 0.023) and  $VO_2$  (p = 0.003). There was not statistically significant interaction between treatment and time. \* $p \le 0.05$ , denote statistically significant differences between groups at the specified time points using the Holm-Sidak test for multiple comparisons.

Accordingly, the present series of experiments supports continuous intraosseous infusion of VP along with low-volume fluid resuscitation using NS for early and sustained hemodynamic stabilization during severe hemorrhagic shock in the battlefield pending evacuation to a medical treatment facility. VP infusion was associated with increases in systemic vascular resistance, reductions in cardiac index, and accentuation of lactic acidosis but reduced the risk of cardiac arrest suggesting a redistribution of blood flow from non-immediately vital territories towards vital organs. NS was associated with preload augmentation resulting in increases in cardiac index and reductions in lactic acid with the combination of VP and NS yielding 100% survival at 72 hours without organ dysfunction. The addition of EPO had no beneficial effect on initial resuscitation and survival contrary to the initial hypothesis. I fact, the data suggested a possible negative effect of EPO on initial resuscitability. The levels of severity; increasing the blood volume removed from 65% to 75%, had minimal detrimental effect on survival that was statistically insignificant suggesting that survival with the combination of VP and NS is feasible despite acute loss of 75% of the blood volume.

We are currently examining the effects of VP and NS as described above in a model of uncontrolled bleeding caused by liver laceration to more closely mimic clinical scenarios in which bleeding in the battlefield is initially uncontrolled and accompanied by tissue injury.

#### KEY RESEARCH ACCOMPLISHMENTS

- The swine model developed by us with funding under award #W81XWH-11-2-0019 was highly effective in reproducing key features of severe hemorrhagic shock allowing assessment of the proposed interventions on initial resuscitation and subsequent survival while exploring the underlying mechanisms.
- Additional software development allowed modeling the blood volume removal pattern to more closely reflect spontaneous bleeding. Such capability can be used in future protocols to model any pattern of blood loss the investigators might want to examine.
- The findings strongly support the use of VP infusion started early after recognition of severe bleeding following by administration of NS in small bolus amounts for hemodynamic stabilization until definitive care is available.

#### REPORTABLE OUTCOMES

#### **PRESENTATIONS**

Targeting Venous Capacitance during Resuscitation from Hemorrhagic Shock. Grand Rounds, Captain James A. Lovell Federal Health Care Center North Chicago, IL, August 2, 2012.

Novel Experimental Approaches to Resuscitation from Hemorrhagic Shock: A Report from the Resuscitation Institute. Department of Medicine Grand Rounds. Rosalind Franklin University of Medicine and Science/The Chicago Medical School, North Chicago, IL, January 9, 2013.

Venous Tone Augmentation with Vasopressin for Hemodynamic Stabilization during Hemorrhagic Shock (lecture, May 31); The Wolf Creek XII Conference. Westin Mission Hills Resort & Spa, Rancho Mirage, CA, May 31, 2013.

#### **ABSTRACTS**

- 1. Borovnik-Lesjak V, Whitehouse K, Baetiong V, Currie B, Radhakrishnan J, Gazmuri RJ. Identification of critical level of blood volume reduction in a swine model of hemorrhagic shock. *Circulation* 2012;126:A12073.
- 2. Gazmuri RJ, Whitehouse K, Borovnik-Lesjak V, Baetiong A, Radhakrishnan J. Vasopressin infusion during severe hemorrhagic shock increases systemic blood flow and markedly improves survival in a swine model. *Circulation* 2012;126:A15905.
- 3. Gazmuri RJ, Whitehouse K, Whittinghill KL, Baetiong A, Radhakrishnan J. Vasopressin with low-volume resuscitation is highly effective for resuscitation and survival from severe hemorrhagic shock. *Circulation* 2013;128:A341.

# **MANUSCRIPTS**

Effects of intraosseous erythropoietin during hemorrhagic shock in swine. Borovnik-Lesjak V; Whitehouse K; Baetiong A; Miao Y; Currie B; Velmurugan S; Radhakrishnan J; Gazmuri RJ (ready for submission).

# **APPLICATIONS**

Pre-Proposal submitted in response to USAMRMC BAA 13-1 entitled "Hemorrhagic Shock Control by Blood Flow Redistribution and Hemostatic Enhancement."

# **CONCLUSION**

Intraosseous infusion of vasopressin from the onset of hemorrhagic shock until the start of blood reinfusion in combination with normal saline (half the amount of blood withdrawn) given at the end of hemorrhage over 30 minutes was highly effective in securing initial resuscitation and 72-hour survival under conditions of severe hemorrhagic shock, when compared with each intervention alone. Erythropoietin did not contribute to resuscitation and 72-hour survival and it might have had an adverse effect. Intraosseous infusion of vasopressin (delivered using a preloaded syringe and a small pump) and low-volume fluid resuscitation are low-skill techniques that could be deployed in far-forward battlefield operations providing extended hemodynamic stability in wounded soldiers pending evacuation to a medical treatment facility.

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- 1. Eastridge BJ, Mabry RL, Seguin P et al. Death on the battlefield (2001-2011): implications for the future of combat casualty care. *J Trauma Acute Care Surg* 2012;73:S431-S437.
- 2. Gazmuri RJ, Whitehouse K, Borovnik-Lesja V, Baetiong A, Radhakrishnan J. Vasopressin infusion during severe hemorrhagic shock increases systemic blood flow and markedly improves survival in a swine model. Circulation 2012;126:A15905.
- 3. Gazmuri RJ, Whitehouse K, Whittinghill KL, Baetiong A, Radhakrishnan J. Vasopressin with low-volume resuscitation is highly effective for resuscitation and survival from severe hemorrhagic shock. *Circulation* 2013;128:A341.

# **APPENDICES**

Curriculum vitae

SUPPORTING DATA

Included in the Body.

#### **CURRICULUM VITAE**

# Raúl J. Gazmuri, MD, PhD, FCCM

# November 20, 2013

#### **PERSONAL**

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Professor (tenure track), Department of Medicine

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#### **EDUCATION**

1991-1994	PhD degree. Department of Physiology and Biophysics. Rosalind Franklin University of Medicine and Science/The Chicago Medical School, North Chicago, IL (June 2, 1994).
1989-1991	Critical Care Medicine training. Department of Medicine, University of Health Sciences/The Chicago Medical School, North Chicago, IL.
1986-1989	Research Fellowship. Department of Medicine, University of Health Sciences/The Chicago Medical School, North Chicago, IL. Laboratory of Max Harry Weil, MD, PhD, Professor and Chairman, Department of Medicine.
1981-1984	Internship and Residency in Internal Medicine. University of Chile, Hospital del Salvador, Santiago.
1974-1980	MD degree. University of Chile School of Medicine.
1962-1973	Primary and High School, Saint Gaspar College, Santiago, Chile.

#### **APPOINTMENTS**

# **Present**

2009-present	Director, Resuscitation	Institute at Rosalind Franklin	University.

- 2002-present Professor (tenure-track). Department of Medicine and Chief, Division of Critical Care Medicine with cross-appointment in the Division of Pulmonary Medicine, Rosalind Franklin University of Medicine and Science/The Chicago Medical School, North Chicago, IL.
- 2009-present Professor. Department of Physiology and Biophysics, Rosalind Franklin University of Medicine and Science/The Chicago Medical School, North Chicago, IL.
- 2000-present Chief, Section of Critical Care Medicine North Chicago VA Medical Center, North Chicago, IL.
- 1998-present ICU Director North Chicago VA Medical Center, North Chicago, IL.

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2002-2009	Associate Professor. Department of Physiology and Biophysics, Rosalind Franklin University of Medicine and Science/The Chicago Medical School, North Chicago, IL.
2002-2006	Site Director (North Chicago VA Medical Center), Critical Care Fellowship Program, Rosalind Franklin University of Medicine and Science/The Chicago Medical School, North Chicago, IL.

- Associate Professor. Department of Medicine, Finch University of Health Sciences/The Chicago Medical School, North Chicago, IL.
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1994-1997	Critical Care Attending. North Chicago VA Medical Center, North Chicago, IL.
1992-1993	Associate Program Director of Critical Care Medicine. University of Health Sciences/The Chicago Medical School. North Chicago, IL.*
1992-1993	Chief, Critical Care Section. Medical Service, North Chicago VA Medical Center, North Chicago, IL.*
1991-1993	Vice-President for Research. Institute of Critical Care Medicine, a Public Educational and Research Foundation. North Chicago, IL, and Palm Springs, CA.*
1991-1992	Attending Physician in Critical Care Medicine, North Chicago VA Medical Center, North Chicago, IL.
1989-1994	Instructor. Department of Medicine. Finch University of Health Sciences/The Chicago Medical School. North Chicago, IL.
1988-1989	Senior Research Fellow and Laboratory Supervisor. Dr. Max Harry Weil's Research Laboratory. University of Health Sciences/The Chicago Medical School.
1986-1989	Research Fellow. Dr. Max Harry Weil's Research Laboratory. University of Health Sciences/The Chicago Medical School.
1984-1986	Staff physician. Center for the Critically Ill, Clínica Las Condes, Santiago, Chile.
1984-1986	Staff physician. Intensive Care Unit, DIPRECA Hospital, Santiago, Chile.

# **COMMITTEE ACTIVITIES (INTRAMURAL)**

2004	Member, Search Committee, Chair Department of Physiology. Rosalind Franklin University of Medicine and Science.
2003-2008	Member, Internal Research Review Panel. Rosalind Franklin University of Medicine and Science.
2001-2006	Member, MD/PhD Combined Degree Admissions Committee. Rosalind Franklin University of Medicine and Science/The Chicago Medical School.
2000-present	Member, Department of Medicine Medical Executive Committee. Rosalind Franklin University of Medicine and Science/The Chicago Medical School.
2001-2003	Member, IACUC. Combined Hines and North Chicago VA committee.
2000-2001	Chair, Research Audit Committee. North Chicago VA Medical Center.
2000-2003	Chair, Department of Medicine Research Committee. Finch University of Health Sciences/The Chicago Medical School.
1998-present	Chair, Critical Care Committee at Captain James A. Lovell Federal Health Care Center.
1998-2011	Chair, CPR Committee at Captain James A. Lovell Federal Health Care Center.
1998-2000	Member of Infectious Control, Transfusion and Blood and Blood Products, and Surgical Case Review Committees. North Chicago VA Medical Center.
1997-1999	Co-Chair, Department of Medicine Research Committee. Finch University of Health Sciences/The Chicago Medical School.

\*Appointments relinquished in 1993 to complete PhD program.

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1997-1998	Chair, Animal Care Committee. Member of Critical Care and CPR, Infectious Control, Transfusion and Blood and Blood Products, Surgical Case Review, and Research and Development Committees. North Chicago VA Medical Center.
1988-1991	Member, Department of Medicine Research Committee. Finch University of Health Sciences/The Chicago Medical School.
1992-1993	Chairman, Critical Care and CPR Committee.  Member, Infectious Disease, Blood Products and Transfusion, Surgical Case Review, and Ethical Advisory Committees. North Chicago VA Medical Center, North Chicago, IL.*

# **COMMITTEE ACTIVITIES (EXTRAMURAL)**

2009-2013	Member, Clinical and Integrative Cardiovascular Science (CICS) Study Section, Center for Scientific Review.
2011-2013	Member, American Heart Association's Advanced Cardiac Life Support (ACLS) Subcommittee.
2006-2011	Member, American Heart Association's Basic Life Support (BLS) Subcommittee (appointed from July 1, 2006 to June 30, 2008; reappointed until June 30, 2011).
2008	Member (temporary), Clinical and Integrative Cardiovascular Science (CICS) NIH Study Section. February 7-8, Washington, DC.
2007	Member (temporary), Myocardial Ischemia and Metabolism (MIM) NIH Study Section. February 22-23, Washington, DC.
2006	Member (temporary), Clinical and Integrative Cardiovascular Science (CICS) NIH Study Section. November 2-3, Washington, DC.
2000-2003	Member, Research Committee, Society of Critical Care Medicine.

MISCELLAN	IEOUS ACTIVITIES
2011-present	Member, International Scientific Committe, A.P.I.C.E. (Anesthesia, Pharmacology, Intensive Care, Emergency).
2009-present	Director, Resuscitation Institute at Rosalind Franklin University.
2004-2008	Medical Director and Founder of the Rosalind Franklin University Life Saving Coalition for improving outcomes after out-of-hospital sudden cardiac arrest. It involves community participation. Donated AEDs to Lake County Sheriff Office [2006] Puerto Rican Society [June 2008].
1998-Present	Software development (for research applications) using LabVIEW, National Instruments.

# SPECIALTY AND SUBSPECIALTY CERTIFICATION

1993	Critical Care Medicine Board (recertified in 2013).
1992	American Board of Internal Medicine (recertified in 2002 and 2012).
1984	Chilean National Examination in Internal Medicine.

# **LICENSURE**

Illinois License, No. 036-078444 1989

1980 Doctor of Medicine and Surgery, Chile

#### PROFESSIONAL SOCIETY MEMBERSHIP

2005-present Honorary Member "Sociedad Decano Lorenzo Sazié." Santiago, Chile

1997-present American Heart Association

1997-present Fellow of the American College of Critical Care Medicine

1990-present Society of Critical Care Medicine, U.S.A.

1990-present American Federation for Medical Research.

1988-1996 American Association for the Advancement of Science.

1986-present Chilean Society of Intensive Care Medicine.

1986-present Medical Society of Santiago, Chile.

1983-present Chilean Medical College.

#### EDITORIAL AND PEER-REVIEW ACTIVITIES

# **Periodicals**

1993-present Reviewer for Critical Care Medicine; Circulation; American Journal of

Physiology; and Journal of Applied Physiology

1996-present Ad-hoc reviewer for Annals of Emergency Medicine; The Lancet; Critical

Care; The Journal of the American College of Cardiology; Expert Opinion on Therapeutic Patents; Resuscitation; and Journal of the American Association

for Laboratory Animal Science; BMC Veterinary Research.

Other

2009-present Associated Editor, Acta Medico-Biotechnica, University of Maribor, Slovenia.

2007-present Member, Scientific Committee, International Seminar on Acute Conditions,

University of Maribor, Maribor, Slovenia.

2006-2009 Member, Faculty of 1000 Medicine, Acute Cardiovascular Problems, Critical

Care & Emergency Medicine.

2003-2012 Abstract reviewer for the AHA Scientific Sessions.

1993-present Occasional reviewer of grant proposals for: FONDECYT (Chilean National

Foundation for Science and Technology); Ministry of Education, Science and

Sport, Republic of Slovenia, and the Austrian Science Fund (FWF).

1991-present Abstract reviewer for the Society of Critical Care Medicine Educational and

Scientific Annual Symposia.

# PROFESSIONAL INVITATIONS

# **Invited Lectures**

1984 [Problems and Complications of Hemodynamic Monitoring] (workshop leader). First Course on Hemodynamic Monitoring. Center for the Critically Ill, Clinica Las Condes, Santiago, Chile.

- [Nutrition in the Patient with Acute Renal Failure]. First Course of Renal Failure in the Critically Ill. Center for the Critically Ill, Clinica Las Condes, Santiago, Chile.
- (a) [Hemodynamic Monitoring], (b) [Nutrition in the Critically Ill], (c) [Airway Management, and Venous Catheterization]. Hospital del Salvador, Santiago, Chile.
- 1986 [Renal Failure in the Critically Ill] (panel). Center for the Critically Ill, Clinica Las Condes, Santiago, Chile.
- 1986 [Septic Shock] (panel). Postgraduate Course. Medical Society of Valparaiso. Valparaiso, Chile.
- 1986 [Nutritional Aspects in Septicemia]. Postgraduate Course in Septic Shock. Medical Society of Valparaiso, Valparaiso, Chile.
- (a) [Hemodynamic Monitoring in Acute Myocardial Infarction], (b) [Problems and Complications in Hemodynamic Monitoring (workshop leader)]. Second Course on Hemodynamic Monitoring. Center for the Critically Ill. Clinica Las Condes, Santiago, Chile.
- 1986 [Pathophysiology, Diagnosis and Treatment of Shock]. Mutual de Seguridad. Santiago, Chile.
- (a) [Clinical Clues in the Diagnosis and Treatment of Shock], (b) [Hemodynamic Monitoring: The Nurse's Role]. Hospital DIPRECA, Santiago, Chile.
- 1986 [Multiple Organ Failure] (panel). Chilean Society of Intensive Medicine. Santiago, Chile.
- [Clinical Application of Electrocardiography] (panel moderator). Chilean Society of Cardiology, Santiago, Chile.
- 1988 [Cardiac Resuscitation by Extracorporeal Pump Oxygenator (ECPO) after Failure of Precordial Compression]. Chicago Cardiology Group, Chicago, IL, U.S.A.
- Extracorporeal Pump Oxygenation (ECPO) During Cardiac Arrest. Fourth Post-Graduate Course "Recent Advances in Anesthesia, Pain, Intensive Care and Emergency. November 21-25. Trieste, Italy.
- 1989 [The Rational Management of Cardiac Arrest. Departments of Anesthesia and Cardiology]. Catholic University of Chile. December 27 and 29. Santiago, Chile.
- (a) Anaphylaxis, (b) Buffers in the Treatment of Acidosis. Twenty-eighth University of Southern California Annual Symposium. Critical Care Medicine. March 12-15. Las Vegas, Nevada, U.S.A.
- 1990 Cardiac Resuscitation by Extracorporeal Circulation. Second International Postgraduate Course in Anesthesia and Intensive Care. June 23. 1990. Alimini Valtur Village, Otranto, (Lecce) Italy.
- (a) [Acute Respiratory Distress Syndrome (panel)], (b) [The Rational Management of Cardiac Arrest], (c) [Cardiac Resuscitation by Extracorporeal Circulation]. Eighth Annual Symposium, Chilean Society of Intensive Care Medicine, November 17-20. Santiago, Chile.
- 1990 [The Rational Management of Cardiac Arrest]. (a) Higueras Hospital, Talcahuano and (b) Sanatorio Alemán, Concepción, November 25; and (c) National Institute of Respiratory Diseases, November 26. Santiago, Chile.
- New Directions for Basic and Advanced Life Support. (a) Medical Grand Rounds, University of Health Sciences/The Chicago Medical School, July 31, North Chicago, IL;

- (b) The Tenth Annual Chicago Critical Care Symposium and Review Course, October 10-13. Chicago, IL; and (c) Cook County Hospital, Critical Care Conference, October 21. Chicago, IL, U.S.A.
- (a) Buffer Agents for the Treatment of Acidosis [panel], (b) Options for Cardiac Resuscitation When Conventional Methods Fail. 30th Annual University of Southern California Annual Symposium on Critical Care Medicine. February 3-7. Las Vegas Nevada, U.S.A.
- Options for Enhancement of Myocardial Perfusion During Cardiac Resuscitation. Seventh Post-Graduate Course. Recent Advances in Anesthesia, Pain, Intensive Care and Emergency. November 25-28. Trieste, Italy.
- Advances in Cardiopulmonary Resuscitation. Slovenian Society of Intensive Medicine. November 30. Ljubljana, Slovenia.
- 1993 Shock What Does it Mean Today? Third Intensive Medicine Symposium. February 26-27. Bremen, Delmenhorst, Germany.
- (a) New Concepts in Cardiac Resuscitation, (b) Use of Capnography in the Critically Ill. Nineteenth Annual Midwest Conference. American Association of Critical Care Nurses. March 24. Arlington Heights, IL, U.S.A.
- (a) Capnography for CPR monitoring, (b) Prehospital CPR by ECPO and Intermittent Aortic Balloon Occlusion. 2nd International Congress on Emergency Medical Assistance. January 25-27. Lyon, France.
- 1994 [Myocardial Dysfunction after Resuscitation from Cardiac Arrest]. DIPRECA hospital. July 8. Santiago, Chile.
- (a) Current Approach to the Treatment of Cardiac Arrest Chain of Survival, (b)
  Options to Restore Myocardial Perfusion. Third International Symposium on Intensive
  Care medicine. September 22-23. Bled, Slovenia.
- (a) [Is Positive Pressure Ventilation Essential During Early CPR?], (b) [Monitoring the Resuscitation Effort], (c) [Advances in Cardiopulmonary Resuscitation]. LXVII Chilean and International Congress of Surgery. November 28 December 1. Santiago, Chile.
- 1996 Cardiopulmonary Resuscitation: The Search for More Effective Options. Medical Grand Rounds. Finch University of Health Sciences/The Chicago Medical School. April 3. North Chicago, IL.
- 1996 Clinical Use of Capnography. Third Joint Meeting of the Slovenian and Croatian Societies of Intensive Care Medicine. May 31 June 1. Otocec, Slovenia.
- Myocardial Effects of Ventricular Fibrillation and Reperfusion in the Isolated Rat Heart. Department of Medicine Quarterly Research Seminar. Finch University of Health Sciences/The Chicago Medical School. September 18. North Chicago, IL.
- Mechanisms of Myocardial Pump Failure After the Global Ischemic Injury of Cardiac Arrest. Department of Microbiology and Immunology. Finch University of Health Sciences/The Chicago Medical School. November 12. North Chicago, IL.
- (a) [The Pathophysiological Bases of Cardiopulmonary Resuscitation]; (b) [Clinical Effectiveness of New Resuscitation Techniques]; (c) [Hemodynamic Guidance During Cardiac Resuscitation]; (d) [Myocardial Protection During Cardiac Arrest]; (e) [Chain of Survival: Community Involvement and Access to Emergency Services]; (f) [Ethical and Legal Aspects of Cardiopulmonary Resuscitation] (panel). First Symposium on

- Cardiopulmonary Resuscitation. Universidad Católica de la Santísima Concepción, Facultad de Ciencias de la Salud. May 7-8. Concepción, Chile.
- Advances in CPR: Medical Grand Rounds. Finch University of Health Sciences/The Chicago Medical School. December 17. North Chicago, IL.
- Resuscitation of the Arrested Heart: Research Activities at North Chicago. Department of Medicine Annual Retreat. Finch University of Health Sciences/Chicago Medical School.
- Options for Myocardial Protection during the Global Ischemic Injury of Cardiac Arrest. Department of Physiology and Biophysics Seminar. Finch University of Health Sciences/The Chicago Medical School. March 25. North Chicago, IL.
- The Effects of Repetitive Electrical Shocks on Post-Resuscitation Myocardial Function. Wolf Creek V Conference. The Institute of Critical Care Medicine. September 8-9. Rancho Mirage, CA.
- Myocardial Protection by Inhibition of Sarcolemmal Sodium Influx. Bioengineering New Resuscitation Therapies for the Treatment of Cardiac Arrest. Emergency Resuscitation Center. November 17-18. Argonne National Laboratory, Argonne, IL.
- 2000 Myocardial Protection by Inhibition of Sarcolemmal Sodium Influx. Naval Dental Research Institute/United States Army Dental Research Detachment/North Chicago VA Medical Center Scientific Colloquium. May 5. Great Lakes Navy Hospital, Great Lakes, IL.
- 2000 Successful Ventricular Defibrillation by Inhibition of the Sodium-Hydrogen Exchanger Isoform-1. Fifth Annual Meeting of Midwest Physiological Society. June 5-6. Finch University of Health Sciences/The Chicago Medical School, North Chicago, IL.
- 2000 NHE-1 Inhibition: A New Treatment Approach for Cardiac Resuscitation. Medical Grand Rounds, June 7, Finch University of Health Sciences/The Chicago Medical School, North Chicago, IL.
- Scientific presentation: Sodium Hydrogen Exchange Inhibitor for Minimizing Post-Resuscitation Myocardial Dysfunction. Panels: Mechanisms and Management of Post-Resuscitation Myocardial Dysfunction; Anti-Arrhythmic Drugs for Facilitating CPR; and Criteria for Discontinuing CPR Efforts. Sixth Wolf Creek Conference on Cardiopulmonary Resuscitation, June 4<sup>th</sup> 7th, The Institute of Critical Care Medicine, Rancho Mirage, CA.
- Beneficial Effects of NHE-1 Inhibition in a Rat Model of Cardiac Arrest. The XVII World Congress of the International Society for Heart Research, July 10th, Winnipeg, Manitoba, Canada.
- Improving Resuscitation from Cardiac Arrest by Inhibition of the Sodium Hydrogen Exchanger Isoform 1. Aventis Pharma, August 16th, Frankfurt, Germany.
- Myocardial Protection during Ventricular Fibrillation by Inhibition of the Sodium-Hydrogen Exchanger Isoform-1. Pulmonary and Critical Care Medicine Grand Rounds, August 24th, University Hospitals of Cleveland and Case Western Reserve University, Cleveland, OH.
- Acute Complications of Pericardial Disease or Injury. February 19th. Acute
  Complications of Diabetes Mellitus. February 20th. An Update on the Clinical
  Presentation and Management of Diabetes Insipidus. February 20th. 40th Anniversary
  Symposium on Critical Care, Trauma, and Emergency Medicine. Keck School of

- Medicine, University of Southern California and The Institute of Critical Care Medicine, February 18 22, Ceasars Palace Hotel, Las Vegas, Nevada.
- 2002 NHE-1 Inhibition: A Potential New Treatment for Resuscitation from Cardiac Arrest. Medical Grand Rounds, April 4, Finch University of Health Sciences/The Chicago Medical School, North Chicago, IL.
- Optimal Timing for Electrical Defibrillation. Resuscitation and Bioengineering Science. Emergency Resuscitation Center. The University of chicago and Argonne National Laboratory, April 18-19, Argonne National Laboratory, Argonne, IL
- NHE-1 Inhibition: A New form of Treatment for Resuscitation from Ventricular Fibrillation. Center for Magnetic Resonance Research, May 8, Evanston, IL
- NHE Inhibition in Cardiac Arrest. Advances in Na-H Exchange Research: From Molecular Regulation to Therapeutic Development. A Satellite Symposium of IUPHAR 2002. The University of Western Ontario, July 12-15, London, Ontario, Canada.
- Abormalities in Water Metabolism (March 19th), CPR tutorial, and New Oprions in CPR (March 20th). Twenty Third International Symposium on Intensive Care and Emergency Medicine. Organized by the Free University of Brussels and the Belgian Society of Intensive Care and Emergency Medicine. Brussels Congress Center, March 18-21, Brussels, Belgium.
- 2003 Reanimación Cardiopulmonar Avanzada (November 6th), Algunas Sugerencias para hacer Investigación en Medicina Intensiva, Sodium-Hydrogen Exchanger Inhibition: A Potencial New Treatment for Cardiac Resuscitation, Augmenting Venous Return during Closed-Chest Resuscitation (November 7th), Mesa Redonda: Aspectos Clave en Paro (November 8th). XXI Congreso Chileno de Medicina Intensiva, November 5-8, Pucón, Chile.
- 2003 Myocardial Dysfunction Following Restoration of Spontaneous Circulation (November 16th). 18th International Symposium on Critical Care Medicine. APICE 2003, November 14-18, Trieste, Italy.
- Experimental and Clinical Advances in Cardiopulmonary Resuscitation. Department of Medicine Grand Rounds, May 20, Rosalind Franklin University of Medicine and Science/The Chicago Medical School, North Chicago, IL
- Minimizing Myocardial Ischemia and Maximizing Myocardial Protection during Resuscitation from Cardiac Arrest (lecture, June 25); Focus on Myocardial Preservation, Reperfusion Injury, Post-Resuscitation Myocardial Dysfunction, and the Role of Buffer Agents during Cardiac Arrest (panelist, June 25); From Basic to Advanced Life Support (panelist, June 26); Revisting the Guidelines (panel cochairman, June 26); Promising Pharmacological Agents for CPR (lecture, June 26); and Technological Innovations to Coordinate and Prioritize Electrical and Mechanical Interventions (panelist, June 26). First Safar-Weil Conference on Cardiopulmonary and Cerebral Resuscitation. June 25-27, Venezia, Italy.
- NHE-1 Inhibition for Resuscitation from Sudden Cardiac Death. Aventis Pharma. June 28, Frankfurt, Germany.
- NHE-1 Inhibition for Resuscitation from Sudden Cardiac Death. Department of Anesthesia, Leopold Franzens University of Innsbruck. June 29, Innsbruck, Austria.
- NHE-1 Inhibition and Coronary Blood Flow during VF are both Determinants of Post-Resuscitation Myocardial Function. Resuscitation Science Symposium (ReSS), grantees

- meeting, co-sponsored by the National Heart, Lung, and Blood Institute and the American Heart Association. November 5-6, New Orleans, LA.
- 2004 Myocardial Preservation, Reperfusion Injury, and Post-Resuscitation Myocardial Dysfunction (lecture, November 13th); Cardiac Arrest (session chair and panelist, November 13th). International Symposium on Critical Care Medicine, 19th Annual Meeting (APICE). November 12-15, Trieste, Italy.
- 2005 Sodium-Hydrogen Exchange (NHE-1) Inhibition: A Promising New Approach to Cardiac Resuscitation. Distinguished Lectures Series, Heart and Stroke Foundation of Ontario Program in Heart Failure and Department of Physiology & Pharmacology, University of Western Ontario, March 7, London, Ontario, Canada.
- 2005 Myocardial Protection during Cardiac Resuscitation by Limiting Sarcolemmal Sodium Entry. National Heart, Lung, and Blood Institute, March 18, Bethesda, MD.
- 2005 Sudden Cardiac Arrest and Mitochondrial Injury: A New Therapeutic Target? Research Symposium, North Chicago VA Medical Center, March 25, North Chicago, IL.
- 2005 Cariporide Enables Hemodynamically more Effective Chest Compression by Leftward Shift of the Flow-Depth Relationship. The 8th Wolf Creek Conference on Cardiopulmonary Resuscitation. June 10-13, The Lodge Hotel at Rancho Mirage, CA.
- Pathophysiology and Management of Myocardial Injury during Cardiopulmonary Resuscitation. 2nd Seminar with International Attendance. Emergency Conditions: Signs, Symptoms, Syndromes, Differential Diagnosis and Management, October 7, Congress Center Habakuk, Maribor, Slovenia.
- 2005 Myocardial Protection by NHE-1 Inhibition. Resuscitation Science RFAs Grantees' Meeting, October 21, Two Rockledge Centre, Bethesda, Maryland.
- 2005 Cardiac Resuscitation (Presentation at Plenary Session). Chest 2005, American College of Chest Physicians, October 21, Montréal, Québec, Canada.
- Basic Science and Pharmacology of Resuscitation (abstract oral session co-moderator). American Heart Association, Scientific Sessions 2005, November 14, Dallas, TX.
- Minimizing Reperfusion Injury in Settings of Myocardial Ischemia (lecture, November 20); Cardiovascular Advances (session co-chair, November 20); CPR Guidelines 2005: What Is New Under The Sun? (panel discussion co-chair, November 20). International Symposium on Critical Care Medicine, 20th Annual Meeting (APICE). November 18-21, Trieste, Italy.
- 2006 Resuscitation from Cardiac Arrest: Are Mitochondria at the Crossroads of Life and Death? Department of Physiology and Biophysics Seminar. Rosalind Franklin University of Medicine and Science. April 20. North Chicago, IL.
- 2006 Importance of Mitochondria during Resuscitation from Cardiac Arrest (April 27); Preclinical Resuscitation Network: A Progress Report (April 28) (self-chosen topics as Program Director). Symposium on Sudden Cardiac Arrest: From Research to Resuscitation, held at the Rosalind Franklin University of Medicine and Science, April 27-29. North Chicago, IL.
- 2006 Guidelines for CPR and Emergency Cardiac Care: Renewed Emphasis on Blood Flow (lecture, May 11); Technological Advances in Resuscitation: A way to Implement Guidelines and Beyond (lecture, May 11); A Practical Approach to Fluid and Electrolyte Management (lecture, May 13). [International Course on Medical Emergencies and

- Critical Ill Patient, organized by a 16-hospital Chilean ICU network]. Hotel Crowne Plaza, May 11-13, Santiago, Chile.
- 2006 Resuscitation 2006. Clínica Las Condes, May 18, Santiago, Chile.
- 2006 Resuscitation 2006. Clínica Dávila, May 19, Santiago, Chile.
- Moderator for Panel Discussion. Advances in Post-Resuscitation Care. Organized by the Emergency Resuscitation Center, the University of Chicago and Argonne National Laboratory. The Gleacher Center, June 10, Chicago, IL.
- Advances in Cardiopulmonary Resuscitation: From Bench to Bedside. Department of Medicine Grand Rounds, July 5, Rosalind Franklin University of Medicine and Science/The Chicago Medical School, North Chicago, IL.
- Novel Targets for Ameliorating Injury Caused by Ischemia and Reperfusion.

  Department of Medicine Research Symposium. August 23, Rosalind Franklin University of Medicine and Science/The Chicago Medical School, North Chicago, IL.
- 2006 Basic CPR Physiology and New Research Directions. October 6, UT Southwestern Medical Center. Children's Medical Center, Dallas, TX.
- 2006 Cardiopulmonary resuscitation: Bridging the Gap between Knowledge and Practice. October 17, Lake County Medical Society, Lincolnshire, IL.
- 2007 Targets for Myocardial Protection during Resuscitation from Cardiac Arrest. Medical Service, Denver VA. February 9, Denver, CO.
- Management of Life Threatening Electrolyte Disorders. March 7th. Recognition and Management of Intrathoracic Crises with Special Reference to Cardiac Tamponade. March 9th. 45th Annual Symposium on Critical Care, Trauma and Emergency Medicine. Keck School of Medicine, University of Southern California, and The Weil Institute of Critical Care Medicine, March 5 9, Ceasars Palace Hotel, Las Vegas, Nevada.
- Activation of the Mitochondrial Apoptotic Pathway without Evidence of Apoptotic Cell Death after Resuscitation from Ventricular Fibrillation (lecture, June 23). Optimizing Post Resuscitation Management II Priorities of Post Resuscitation Care, Do we as yet have enough Evidence to Justify a Unified Algorithm? (panel, June 23). Optimizing Transitions from Laboratory to Clinic (panel, June 23). The Wolf Creek IX Conference. June 21-24, The Renaisance Esmeralda Resort & Spa, Indian Wells, CA.
- 2007 In-Hospital Resuscitation (forum). Clínica Dávila, August 6, Santiago, Chile.
- 2007 Practical Concepts in Fluid Management. North Chicago VA Grand Rounds, September 4, North Chicago, IL.
- 2007 Myocardial Effects of Cardiac Arrest and Resuscitation with Especial Reference to Mitochondrial Injury (lecture). University of Maribor, Faculty of Medicine, October 3, Maribor, Slovenia.
- 2007 Myocardial Effects of Cardiac Arrest and Resuscitation with Especial Reference to Mitochondrial Injury (lecture, October 5); Critical Care Knowledge in Emergency Medicine. Emergency Conditions: Signs, Symptoms, Syndromes, Differential Diagnosis and Management, October 4-6, Congress Center Habakuk, Maribor, Slovenia.
- 2008 Sodium-Hydrogen Exchange Inhibition during Resuscitation: Saving Myocardium by Protecting Mitochondria? (lecture, hospital, April 23). Perspectives on How to Improve

- Resuscitation Outcomes (lecture, outpatient center, April 23; medical school and school of nursing, April 24). Rounds in various ICUs (April 23). Maribor, Slovenia.
- 2008 Carbon Dioxide during Resuscitation: A Gas to Monitor and a Molecule to Understand; Sodium Hydrogen Exchange Inhibition and other Pharmacological Interventions during Resuscitation: Saving Myocardium by Protecting Mitochondria? (lectures, April 28). Physio-Control. Perspectives on Improving Resuscitation Outcomes. Seattle Resuscitation Rounds. (lecture, April 29). Harborview Medical Center. Seattle, Washington.
- Conference: Myocardial protection after CPR; Practical Lessons Learned on How to Setup a Research Laboratory (May 8); A Practical Approach to Fluid Management (May 9); Cardiopulmonary Resuscitation: Bridging the GAP Between Knowledge and Practice (May 10). Physio-Control. XIII Brazilian Congress of Intensive Care Medicine and 1st International Symposium of the World Federation of Intensive and Critical Care Medicine. Salvador City, Bahia, Brazil, May 6 10.
- 2008 [Clinical Research Priorities in Cardiopulmonary Resuscitation, June 9]. [XLIII National Congress of the SEMICYUC and XXXIV National Congress of the SEEIUC]. Valencia, Spain, June 9-11.
- 2008 Advances in Cardiopulmonary Resuscitation (August 28). Clínica Dávila, Santiago, Chile.
- 2008 Erythropoietin for Cardiac Resuscitation. Department of Medicine Grand Rounds, Rosalind Franklin University of Medicine and Science/The Chicago Medical School. North Chicago, IL, July 30.
- Targeting Mitochondria for Resuscitation from Cardiac Arrest. Department of Medicine Grand Rounds, Rosalind Franklin University of Medicine and Science/The Chicago Medical School. North Chicago, IL, September 17.
- 2008 [Scientific Knowledge Gap on Resuscitation and Priorities in Clinical Resuscitation for ILCOR 2010, October 3]. [First Canary Conference on Resuscitation], Santa Cruz de Tenerife, Canary Islands, Spain, October 2-3.
- Enhancing the Hemodynamic Efficacy of Chest Compression by Preserving Mitochondrial Bioenergetic Function. Cardiovascular Research Center's Visiting Faculty Seminar Series, Mount Sinai School of Medicine, New York, NY, January 16.
- 2009 [Latest Advances in the Chain of Survival] and [Improving the Hemodynamic Efficacy of Chest Compression by Maintaining Mitochondrial Function] (lectures). [II Conference on Resuscitation. Cerebral Survival, SESCAM Office of Urgencies and Emergencies], Toledo, Spain, April 29.
- 2009 Erythropoietin Facilitates the Return of Spontaneous Circulation and Survival in Victims of Out-of-Hospital Cardiac Arrest. Hines/North Chicago Research Day, Hines, IL, May 6.
- 2009 Improving CPR Hemodynamics and Outcomes by Preserving Left Ventricular Distensibility. Department of Medicine Grand Rounds. Rosalind Franklin University of Medicine and Science/The Chicago Medical School, North Chicago, IL, August 19.
- 2009 Key Points for Improving the Outcome of Out-of-Hospital Cardiac Arrest (lecture, August 31), Inotropic Options for Post-Resuscitation Myocardial Dysfunction (lecture, August 31), CPR: From Flying Blind to Flying Right (lecture, September 1), Cardiopulmonary Resuscitation (moderator, August 31). 10th Congress of the World

- Federation of Societies of Intensive and Critical Care Medicine (WFICCM), Fortezza de Baso, Florence, Italy, August 28 September 1.
- NHE-1 Inhibitors and Erythropoietin for Maintaining Myocardial Function during Cardiopulmonary Resuscitation (lecture October 16), Cardiopulmonary Resuscitation: From Flying Blind to Flying Right (lecture October 16). Acute Conditions: Signs, Symptoms, Syndromes, Differential Diagnosis and Management. Congressional Center Habakuk, Maribor, Slovenia, October 15-17.
- 2009 Medical Emergency Response System (MERS) at the North Chicago VAMC (Grand Rounds). Advances in Medicine, Surgery, and Psychiatry, Department of Veterans Affairs North Chicago and Rosalind Franklin University of Medicine and Science. North Chicago, IL, November 3.
- 2010 Advances in Resuscitation. Department of Medicine Grand Rounds. Rosalind Franklin University of Medicine and Science/The Chicago Medical School, North Chicago, IL, February 10.
- 2010 Resuscitative Medicine: Improving CPR Hemodynamics and Outcomes by Preserving Left Ventricular Distensibility. The Heart Institute for Children Research Club. Hope Children's Hospital, Oak Lawn, IL, March 5.
- 2010 Key Factors for Improving Outcome after Sudden Cardiac Arrest (lecture, March 19); Improving CPR Hemodynamics by Maintaining Left Ventricular Distensibility (lecture, March 19); An Elective in Critical Care Medicine for Second-Year Students (panel New Directions in Health Professions Education, March 19). 65th Annual Medwest Clinical Conference & Technology Summit. Westing Lombard Yorktown Center, Lombard, IL, March 18-19.
- 2010 Protecting Mitochondrial Bioenergetic Function during Resuscitation from Cardiac Arrest. Kick-Off Speaker Session, Hines/North Chicago VA Research Poster Day. Hines VA, Hines, IL, April 29.
- 2010 Protecting Mitochondrial Bioenergetic Function. First Maribor Resuscitation Summit: On The Future of CPR." Maribor, Slovenia, September 9-11.
- Pathophysiology and Severity of Post-Resuscitation Myocardial Dysfunction (lecture); ACLS Recall on Skills and Pharmacological Agents (panel); chairman for three sessions on CPR topics. APICE 23rd International Symposium on Critical Care Medicine, Catania, Italy, November 5-7.
- [Latest Recommendations in Resuscitation] and [Science of Resuscitation] (lectures). [III Conference on Advances in Resuscitation. New Recommendations and New Futures, SESCAM Office of Urgencies and Emergencies], Toledo, Spain, December 16.
- Recognition and Treatment of Physiological Deterioration and Other Life-Threatening Emergencies through Implementation of a Rapid Response System at the Captain James A. Lovell Federal Health Care Center. Grand Rounds, Captain James A. Lovell Federal Health Care. Center North Chicago, IL, May 5.
- Minimal Hemodynamic effects of Increasing Respiratory Rate and Volume during CPR but large Impact on P<sub>ET</sub>CO<sub>2</sub> (lecture, June 11). Optimizing the Efficacy of Circulatory Support during CPR (panel, June 11). Optimizing Transitions from Laboratory to Clinic (panel, June 11). The Wolf Creek XI Conference. Westin Mission Hills Resort & Spa, Rancho Mirage, CA, June 9-12.
- 2011 Prevention of in Hospital Cardiac Arrest by Early Recognition and Treatment of Physiological Deterioration. Department of Medicine Grand Rounds. Rosalind Franklin

- University of Medicine and Science/The Chicago Medical School, North Chicago, IL, September 14.
- 2011 Cardiac arrest From the Lab to Clinical Trials (lecture, December 11); A Giant of Critical Care Prof. Max Harry Weil (panel, December 11). APICE Masterclass. Annual International Meeting Mediterranean School of Intensive and Critical Care Medicine. Catania University, Catania, Italy, December 9 11.
- Targeting Venous Capacitance during Resuscitation from Hemorrhagic Shock. Grand Rounds, Captain James A. Lovell Federal Health Care Center North Chicago, IL, August 2.
- Targeting Mitochondria during CPR (lecture, September 9); State-of-the-Art in Critical Care (panel, September 8); Quality of CPR (panel, September 8); Post-Resuscitation Care (panel, September 9). Weil Conference on Cardiac Arrest, Shock and Trauma. Istituto Mario Negri, Milan, Italy, September 8–9.
- 2012 [2010 Guidelines for Cardiopulmonary Resuscitation. Highlights] (lecture, November 12); [Physiological Principles of Cardiopulmonary Resuscitation] (lecture, November 12); Mitochondrial Protection during Cardiac Resuscitation (lecture, November 14); [Key Factors for Successful Resuscitation (lecture, November 14). XXXI World Congress of Internal Medicine. Santiago, Chile, November 11-15.
- Novel Experimental Approaches to Resuscitation from Hemorrhagic Shock: A Report from the Resuscitation Institute. Department of Medicine Grand Rounds. Rosalind Franklin University of Medicine and Science/The Chicago Medical School, North Chicago, IL, January 9.
- New Developments in Cardiopulmonary Resuscitation (leture). University of Maribor, Faculty of Medicine. University of Maribor, Slovenia, March 22.
- Current Determinant of Successful Resuscitation and Suggestions for Future Research (lecture, March 22). Acute Conditions (5<sup>th</sup>): Signs, Symptoms, Syndromes, Differential Diagnosis and Management. Congressional Center Habakuk, Maribor, Slovenia, March 21-23.
- Venous Tone Augmentation with Vasopressin for Hemodynamic Stabilization during Hemorrhagic Shock (lecture, May 31); Estrogen Fails to Facilitate Resuscitation from VF in Male Rats (June 1). Optimizing Monitoring and Management of Circulatory Shock: Goal Directed Therapy (panel, May 31); After More than 50 Years of Modern CPR, do we Really Save More Lives? What are the Major Limiting Factors that Prevent Us to Do So? (panel, May 31); Optimizing the Efficacy of Circulatory Support during CPR: The role of Mechanical Devices (panel, June 1); Optimizing Transitions from Laboratory to Clinical Practice (panel, June 1); Optimizing Post-Resuscitation Management: Priorities of Post-Resuscitation Care. The Wolf Creek XII Conference. Westin Mission Hills Resort & Spa, Rancho Mirage, CA, May 30–June 2.
- 2013 Myocardial Effects of Protecting Mitochondrial Function during Resuscitation from Cardiac Arrest (lecture, September 20). American Heart Association 2013 Chicago Research Network Symposium: Transgenic Approaches to Cardiovascular Disease: Past, Present, and Future. Loyola University Chicago, Maywood,IL, September 20.

#### **Other Invitations**

To attend the International Guidelines 2000 Conference on CPR and ECC. American Heart Association. February 6-8. Dallas, TX.

- To serve as expert reviewer for the International Liaison Committee of Resuscitation's (ILCOR) 2005 International Consensus on Emergence Cardiovascular Care (ECC) and Cardiopulmonary Resuscitation (CPR) Science with Treatment Recommendations Conference (C2005), reviewing the topic "Is CPR before defibrillation safe, effective, and feasible?"
- 2003 To serve as section editor for the June 2003 and June 2004 issues of *Current Opinion in Critical Care* dedicated to Cardiopulmonary Resuscitation.
- To serve as invited editor for the November 2005 issues of *Clinic@s de Medicina Crítica* dedicated to Cardiopulmonary Resuscitation.
- 2005 To co-moderate a session on oral presentations on resuscitation topics at the 2005 annual Scientific Sessions of the AHA.
- 2006 To join the APICE 2006 International Scientific Committee.
- To co-moderate a session on oral presentations on resuscitation topics at the 2007 annual Scientific Sessions of the AHA and to co-moderate a poster discussion at the 2007 Resuscitation Science Symposium.
- 2008 Worksheet presenter at ILCOR meeting (November 5); Co-moderator of a poster session at the Resuscitation Science Symposium (November 9); Co-moderator of an abstract presentation session at AHA Scientific Sessions (November 10).
- To serve as expert reviewer for the International Liaison Committee of Resuscitation's (ILCOR) 2010 International Consensus on Emergence Cardiovascular Care (ECC) and Cardiopulmonary Resuscitation (CPR) Science with Treatment Recommendations Conference (C2010).

# **COURSES ORGANIZED**

- [Second Course of Hemodynamic Monitoring (Coordinator)]. Center for the Critically Ill, Clinica Las Condes, Santiago, Chile.
- 1986 [Diagnosis of Emergencies in Intensive Medicine (Director)]. DIPRECA Hospital, Santiago, Chile.
- Sudden Cardiac Death: From Cell Physiology to Clinical Resuscitation. A symposium jointly sponsored by the Department of Medicine and the Department of Physiology and Biophysics and by the Medical Service at the North Chicago VA Medical Center. October 23. North Chicago, IL. Role: Program Director and Speaker. Invited speakers included several nationally recognized investigators.
- Symposium on Sudden Cardiac Arrest: From Research to Resuscitation, held at the Rosalind Franklin University of Medicine and Science, April 27-29. Program Director, Speaker, and Moderator. Invited speaker included 14 internationally recognized leaders in CPR.
- 2006 ICU Elective for 2<sup>nd</sup> year medical students (MMED 699 Critical Care Medicine). Fall quarter 12 sessions, 2 hours each.
- First Maribor Resuscitation Summit: On The Future of CPR." Maribor, Slovenia, September 9-11. Co-Organizer with Dr. Štefek Grmec. Invited speaker included 16 internationally recognized leaders in CPR.

# HOST OF RECOGNIZED SCIENTISTS AT DEPARTMENTAL LEVEL

- Morris Karmazyn, PhD, Canada Research Chair in Experimental Cardiology; Director, Heart and Stroke Foundation of Ontario; Professor of Physiology and Pharmacology, University of Western Ontario, Canada. *Mediation of Ischemic Myocardial Injury, Hypertrophy and Heart Failure by Sodium-Hydrogen Exchange*. Rosalind Franklin University of Medicine and Science, May 26.
- 2008 Roger J. Hajjar, MD, Professor of Medicine and Director of the Cardiovascular Research Center at Mount Sinai Medical Center, New York, NY. *Targeting Calcium Cycling in Heart Failure*. Department of Physiology & Biophysics seminar series. Rosalind Franklin University of Medicine and Science, March 27.

# HOST OF THE RESUSCITATION INSTITUTE LECTURE SERIES

- Tom P. Aufderheide, MD, FACEP, FACC, FAHA; Professor of Emergency Medicine; Associate Chair of Research Affairs; Director, Adult Translational Research Unit; Department of Emergency Medicine; Medical College of Wisconsin. *Harnessing the Thoracic Pump to Treat Patients in Cardiac Arrest: From 1960 to the Present* (March 12).
- Terry L. Vanden Hoek, MD, FACEP; Professor and Chair Department of Emergency University of Illinois at Chicago. *Cardiopulmonary Resuscitation in 2012: New Opportunities for Tranlational Science* (June 8).
- Morris Karmazyn, PhD; Canada Research Chair in Experimental Cardiology; Professor, Department of Physiology and Pharmacology; University of Western Ontario, Canada. The Myocardial Sodium-Hydrogen Exchanger and its Role in Heart Disease: From Basic Research to Clinical (Mis)Trials (November 29).
- Wanchun Tang MD, MCCM, FCCP, FAHA Professor, President, and Chief Scientific Officer of the Weil Institute of Critical Care Medicine; Clinical Professor, Keck School of Medicine, University of Southern California; Professor of Emergency Medicine, UC San Diego School of Medicine. *Recent Advances in Cardiopulmonary Resuscitation* (August 21).

# **HONORS AND AWARDS (First or Senior Author)**

- Acknowledgment to the best scientific project. Norgine Laboratory Prize. XXI Chilean Congress of Cardiology and Cardiovascular Surgery. December, Viña del Mar, Chile. [Arterial and venous prostagladin E<sub>2</sub> levels in patients with valvular disease]. Guarda E, Zamorano B, **Gazmuri RJ**, Escobar E.
- Best Poster Prize. Annual meeting of Swiss Society of Intensive Care Medicine. September, Montreaux, Switzerland. [Non-invasive hemodynamic monitoring during cardiopulmonary resuscitation using capnography]. von Planta M, Weil MH, **Gazmuri RJ**.
- First Prize in Basic Sciences. 18<sup>th</sup> Annual Scientific Meeting of The Society of Critical Care Medicine. June, New Orleans, Louisiana, U.S.A. Myocardial depressant effect of carbon dioxide in the intact rat heart. Wanchun T, **Gazmuri RJ**, Weil MH, Kette F, Bisera J, Duggal C, Rackow EC.
- Honorary Member, Chilean Society of Surgeons. December 1. Santiago Chile.

- Poster award winner at the Annual Loyola/Hines Research Poster Session. April 4, Hines, IL. Left ventricular diastolic dysfunction after resuscitation from cardiac arrest. **Gazmuri RJ**, Berkowitz M, Cajigas H, Hawkins K.
- Emergency Medicine Specialty Award recipient for 2001. Society of Critical Care Medicine 30<sup>th</sup> Educational and Scientific Symposium, February 13, San Francisco, CA. Transtracheal Oxygenation during Cardiac Arrest Maintains its Effectiveness Despite Increased Duration of Chest Compression. Ayoub IM, **Gazmuri RJ**, Brown DJ, Obayashi DY.
- Research Citation (one of 30 semi-finalists). Society of Critical Care Medicine 30<sup>th</sup> Educational and Scientific Symposium, February 13, San Francisco, CA. Cariporide Facilitates Successful Defibrillation and Improves Post-Resuscitation Hemodynamic Function after Prolonged Untreated VF. **Gazmuri RJ**, Ayoub IM, Kolarova J, Damera M.
- Research Citation (one of 6 finalists). Society of Critical Care Medicine 31<sup>st</sup> Critical Care Congress, January 29, San Diego, CA. Successful Treatment of Prolonged Untreated Ventricular Fibrillation Requires Chest Compression before Electrical Shocks. Kolarova J, Yi Zhong, Ayoub IM, **Gazmuri RJ**.
- Emergency Medicine Specialty Award recipient for 2003. Society of Critical Care Medicine 32<sup>nd</sup> Critical Care Congress, February 1, San Antonio, TX. NHE-1 Inhibition Enhances the Pressor Responses to Epinephrine and Vasopressin during Chest Compression. Kolarova J, Yi Z, Ayoub IM, **Gazmuri RJ**.
- Research Citation (one of 5 finalists). Society of Critical Care Medicine 32<sup>nd</sup> Critical Care Congress, February 1, San Antonio, TX. Inhibition of the sodium-hydrogen exchanger isoform-1 (NHE-1) ameliorates reperfusion arrhythmias and prevents episodes of recurrent VF. Ayoub IM, Kolarova J, Yi Z, Maldonado MA, **Gazmuri RJ**.
- Research Citation. Society of Critical Care Medicine 34<sup>th</sup> Critical Care Congress, January 18, Phoenix, AZ. Zoniporide ameliorates post-resuscitation myocardial dysfunction by flow independent mechanisms. Ayoub IM, Kolarova JD, Radhakrishnan J, Wang S, **Gazmuri RJ**.
- Research Citation. Society of Critical Care Medicine 34<sup>th</sup> Critical Care Congress, January 18, Phoenix, AZ. Zoniporide ameliorates intramyocardial lactate increases during resuscitation from ventricular fibrillation. Radhakrishnan J, Wang S, Ayoub IM, Kolarova JD, **Gazmuri RJ**.
- Emergency Medicine Specialty Award. Society of Critical Care Medicine, 35<sup>th</sup> Critical Care Congress, January 10, San Francisco, CA. Cytochrome *c*: a novel biomarker for predicting survival after cardiac resuscitation. Ayoub IM, Wang S, Radhakrishnan J, Kolarova JD, Lyu T, Ryan C, **Gazmuri RJ**.
- Resuscitation/Critical Care Best Abstract Award presented by the American Heart Association's Council on Cardiopulmonary, Perioperative and Critical Care. American Heart Association Scientific Sessions 2008. November 11, New Orleans, Louisiana. Erythropoietin facilitates the return of spontaneous circulation and subsequent ICU admission in victims of out-of-hospital cardiac arrest. Grmec S, Strnad M, Kupnik D, Sinkovic A, Marc G, Gazmuri RJ.
- Cournand and Comroe Young Investigator Prize in Cardiopulmonary and Critical Care (one of 5 finalists), Council on Cardiopulmonary, Critical Care, Perioperative and Resuscitation. American Heart Association Scientific Sessions 2009. November 17, Orlando, Florida. The mitochondrial permeability transition pore opens during

- ventricular fibrillation in a rat model of closed chest resuscitation. Ayoub IM, Radhakrishnan J, Upadhyaya MP, **Gazmuri RJ**.
- Second Prize Poster Winner at the Annual Hines/North Chicago VA Research Day. April 29, Edward Hines, Jr VA Hospital, Hines, IL. Preservation of left ventricular myocardial distensibility using a novel sodium-hydrogen exchanger isoform-1 inhibitor markedly enhances the hemodynamic efficacy of chest compression. Ayoub IM, Upadhyaya MP, Radhakrishnan J, **Gazmuri RJ**.
- Young Investigator Award. American Heart Association Resuscitation Science Symposium 2011. Erythropoietin enables mitochondria to preserve oxidative phosphorylation under conditions of calcium overload and oxidative stress. Radhakrishnan J, Ng M, Edelheit A, **Gazmuri RJ**.

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#### **BOOK CHAPTERS**

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- 2. Shah D, Weil MH, **Gazmuri RJ**, Rackow EC. O<sub>2</sub> Transport in Low Flow States of Circulatory Shock and Cardiac Arrest. <u>In</u>: Reinhart K,Eyrick K (eds). *Clinical Aspects of O<sub>2</sub> Transport and Tissue Oxygenation*. Springer-Verlag, Heidelberg, Germany 1989:445-461.
- 3. Weil MH, **Gazmuri RJ**, Rackow EC. Issues in CPR. <u>In</u>: Fuhrman BP, Shoemaker W (eds). *Critical Care: State of the Art*. The Society of Critical Care Medicine, Fullerton, California, U.S.A. 1989;10: 427-63.
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# PEER REVIEWED ORIGINAL SCIENTIFIC ARTICLES

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- and controlled coronary perfusion by extracorporeal circulation. *Heart Rhythm* 2013;10:S261.
- 152. Borovnik-Lesjak V, Whitehouse K, Baetiong A, Radhakrishnan J, Kaufman CL, **Gazmuri RJ**. Changes in AMSA during controlled coronary perfusion by extracorporeal circulation in a swine model of ventricular fibrillation. *Circulation* 2013;128:A73
- 153. Velmurugan S, Radhakrishnan J, Baetiong A, **Gazmuri RJ**. Marked attenuation of cell injury by substrate availability during ischemia. *Circulation* 2013;128:A340.
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- 155. Whittinghill KL, Miao Y, Radhakrishnan J, **Gazmuri RJ**. Vasodilatory effect of ginseng after resuscitation from cardiac arrest. *Circulation* 2013;128:A147.
- 156. **Gazmuri RJ**, Whitehouse K, Whittinghill KL, Baetiong A, Radhakrishnan J. Vasopressin with low-volume resuscitation is highly effective for resuscitation and survival from severe hemorrhagic shock. *Circulation* 2013;128:A341.

# **PHD THESIS**

**Gazmuri RJ**. Myocardial Function after Resuscitation From Cardiac Arrest. Finch University of Health Sciences/The Chicago Medical School. Department of Physiology and Biophysics, North Chicago, IL 1994.

# THESIS ADVISOR (PHYSICAN ASSISTANT PROGRAM AT FINCH UNIVERSITY OF HEALTH SCIENCES)

Asbell C and Welsh T. In Hospital Resuscitation: Factors Influencing Outcome. North Chicago, IL 1999.

# PHD THESIS MENTOR (DEPARTMENT OF PHYSIOLOGY & BIOPHYSICS AT ROSALIND FRANKLIN UNIVERSITY OF MEDICINE AND SCIENCE)

Iyad M. Ayoub, MS. Mitochondrial Injury during Resuscitation from Cardiac Arrest. North Chicago, IL 2009.

#### **PATENTS**

#### FACILITATION OF RESUSCITATION FROM CARDIAC ARREST BY ERYTHROPOIETIN

Patent No.: US 8,067,36 B2 Date of Patent: Nov 29, 2011 Inventor: Raúl J.Gazmuri Assignee: Rosalind Franklin University of Medicine and Science.

CIRCULATING CYTOCHROME C AS BIOMARKER OF REPERFUSION INJURY AFTER WHOLE BODY ISCHEMIA AND PREDICTOR OF SURVIVAL

Provisional application No. 60/937728 filed on June 29, 2007 (112461.00057) Status, pending.

## **RESEARCH GRANTS (PAST/ACTIVE)**

## PEER REVIEWED

## **Past**

1981-1983 "The Role of Prostaglandins in the Cardiac Adaptation to Overload."

Source: Libraries and Research Department, University of Chile. #1156-8114.

Role: Research Assistant.

Principal Investigator: Edgardo Escobar, MD

1984-1985 Project: The Role of Prostaglandins in the Development of Cardiac

Failure

Source: Libraries and Research Department, University of Chile. #M18761-8413.

Role: Research Associate.

Principal Investigator: Edgardo Escobar, MD

1987-1994 Project: Myocardial Acid-Base Changes During CPR

Source: National Institutes of Health, NHLBI #1R01 HL39148-01 and Competing

Continuation #1Ro1 HL39148-04A2.

Annual Direct Costs: \$82,000 to approximately \$200,000.

Role: <u>Co-Investigator</u>.

Principal Investigator: Max Harry Weil, MD, PhD

1989-1991 Project: Extracorporeal Pump Resuscitation After Prolonged Arrest

Source: American Heart Association, #890912.

Annual Direct Costs: \$29,900. Role: Collaborating Investigator.

Principal Investigator: Max Harry Weil, MD, PhD

1989-1991 Project: Resuscitation after Prolonged Cardiac Arrest

Source: National Institutes of Health, NHLBI #1RO1 HL42590-01.

Annual Direct Costs: \$148,986.

Role: Co-Investigator.

Principal Investigator: Max Harry Weil, MD, PhD

1998-2000 Project: Myocardial Protection During Cardiac Arrest

Source: VA Merit Review.

Annual Direct Cost: \$105,800 to \$109,100.

Role: Principal Investigator.

2001-2004 Project: Myocardial Protection During Ventricular Fibrillation

Source: VA Merit Review.

Annual Direct Cost: \$150,000 to \$225,500.

Role: Principal Investigator.

2002-2006 Project: Myocardial Protection by NHE-1 Inhibition

RFA: Basic Research to Improve Cardiopulmonary and Neurological Outcomes.

Source: NHLBI, 1 R01 HL71728-01.

Annual Direct Cost: \$175,000 to \$225,000.

Role: Principal Investigator.

2005-2006 Project: Evaluation of a New Technique for Controlled Manual

Ventilation

RFA: SBIR/STTR - Technologies for monitoring and performing resuscitation.

Source: NIH.

Annual Direct Cost: \$100,000.

Role: Collaborator.

2006-2007 Project: Myocardial Protection by GSK-3β Inhibition during

Resuscitation from Cardiac Arrest Internal Funding Program (pilot grant).

Source: Rosalind Franklin University of Medicine and Science.

Annual Direct Cost: \$25,000. Role: <u>Principal Investigator</u>.

2006-2007 Project: Bridge Funds

Sponsor: Rosalind Franklin University of Medicine and Science

Total Cost: \$50,000

Role: Principal Investigator.

Active

2010-2014 Project: Myocardial Effects of Erythropoietin during Resuscitation

from Cardiac Arrest Source: VA Merit Review. Annual Direct Cost: \$150,000 Role: <u>Principal Investigator</u>.

2010-2013 Project: Intraosseous Erythropoietin for Acute Tissue Protection in

**Battlefield Casualties Suffering Hypovolemic Shock** 

Source: Defense Medical Research and Development Program (DMRDP), Applied Research and Technology Development Award (ARADTA).

Annual Direct Cost: \$400,000 Role: Principal Investigator.

2012-2013 Project: Plasma Cytochrome c as Biomarker of Traumatic Injury and

**Predictor of Outcome** 

Source: Chicago Medical School and Advocate Lutheran General Hospital

Translational Research Pilot Grant Program Annual Direct Cost (RFUMS): \$70,250

Role: Co-Principal Investigator.

NON-PEER REVIEWED (INVESTIGATOR INITIATED)

1987-1988 Project: Effects of an Extracorporeal Pump Oxygenator in Myocardial

Resuscitability during Experimental Cardiac Arrest

Source: Baxter (American Bentley). Annual Direct Costs: \$82,000.

Principal Investigator: Max Harry Weil, MD, PhD

Role: Co-Investigator.

1995-1997 Project: **Start-up funds** 

Source: Finch University of Health Sciences/The Chicago Medical School.

Total Cost: \$20,000. Role: <u>Principal Investigator</u>.

2000-2001 Project: Bridge funds

Source: Finch University of Health Sciences/The Chicago Medical School.

Total Cost: \$20,000. Role: <u>Principal Investigator</u>.

2004-2005 Project: Myocardial Protection during Ventricular Fibrillation by NHE-

1 Inhibition Using Cariporide: Studies in a Porcine Model

Sponsor: Aventis Pharma, Frankfurt, Germany

Total Cost: \$40,000. Role: <u>Principal Investigator</u>.

Project: Investigation of an NHE-1 Inhibitor for Resuscitation from 2008-2009

Cardiac Arrest

Sponsor: Boehringer Ingelheim Pharmaceuticals, Inc.

Total Cost: \$50,000.

Role: Principal Investigator.

**Project: Volume-Controlled Manual Ventilation during Resuscitation** 2009-2010

from Cardiac Arrest

Sponsor: Dessinier Corporation

Total Cost: \$120.000 Role: Principal Investigator.

2009-2010 Project: Vitamin-C Preserves Myocardial Distensibility during

**Resuscitation from Cardiac Arrest** 

Sponsor: Zdravstveni Dom. Dr. Adolfa Drolca, Maribor, Slovenia

Total Cost: \$20,000

Role: Principal Investigator.

Project: Mechanisms of Cytochrome c Release during Resuscitation – 2010-2010

Adeno-Associated Viral Gene Transfer (Pilot Grant)

Sponsor: Gift for cardiovascular research administered by the Hines VA Research

Service.

Total Cost: \$40,000

Role: Principal Investigator.

NON-PEER REVIEWED (SPONSORED CLINICAL TRIALS)

1999-2000 Project: Antithrombin III in Patients with Severe Sepsis. A

Multinational, Double-Blind, Randomized, Placebo-Controlled Phase-

III Study

Protocol No. BI51.071/7MN-303SE Sponsor: Centeon Pharma GmbH.

Sponsor Representative: PPD Pharmaco International. Role: Principal Investigator at North Chicago VA.

Project: Linezolid (PNU-100766) in the Treatment of Patients with 1999-2000

Nosocomial Pneumonia: A Double-Blind, Randomized, Comparator-

**Controlled Study** 

Protocol No. M/1260/0048 Sponsor: Pharmacia & Upjohn.

Role: Sub-Investigator at North Chicago VA. Principal Investigator: Frank A. Maldonado, MD.

Project: A Randomized, Double-Blind, Double-Dummy, Multicenter 2000-2001

Study to Compare the Efficacy and Safety of Intravenous

Gemifloxacin (SB-265805) Followed by Oral Gemifloxacin Versus Intravenous Lovofloxacin Followed by Oral Levlfloxacin in the Treatment of Hospitalized Adult Patients with Community-Acquired

Pneumonia (CAP)

Sponsor: SmithKine Beecham.

Role: Principal Investigator at North Chicago VA.

2000-2001 Project: A Randomized, Double-Blind, Double-Dummy, Multicenter,

Parallel Group Study to Assess the Efficacy and Safety and Tolerability of Intravenous Gemifloxacin Followed by Oral Gemifloxacin versus Intravenous Ceftriaxone Followed by Oral Levofloxacin in the Treatment of Nosocomial Pneumonia (NP) in Adults at Low Risk for Infection with Pseudomonas Aeroginosa

Sponsor: SmithKine Beecham.

Role: <u>Sub-Investigator</u>.

Principal Investigator: Frank A. Maldonado, MD.

2000-2001 Project: A Randomized, Double-Blind, Phase III, Comparative Study of

Cidecin™ (Daptomycin)to Rocephin® (Ceftriaxone) in the Treatment of Moderate to Severe Community-Acquired Acute Bacterial

Pneumonia Due to S. Pneumoniae

Sponsor: Cubist Pharmaceuticals

Role: Sub-Investigator.

Principal Investigator: Ashok M. Fulambarker, MD.

2008-2010 Project: CV185030: A Phase III, Active (Warfarin) Controlled,

Randomized, Double-Blind, Parallel Arm Study to Evaluate Efficacy and Safety of Apixaban in Preventing Stroke and Systemic Embolism in Subjects with Nonvalvular Atrial Fibrillation (ARISTOTLE)

Sponsor: Bristol Myers Squibb.

Total Cost: \$415,245 (target 19 patients).

Role: Principal Investigator at the North Chicago VA Medical Center.

2008-2010 Project: A Multi-center, Randomized, Double-Blind, Placebo-

Controlled Study to Evaluate the Safety and Efficacy of SCH 530348 in Addition to Standard of Care in Subjects With Acute Coronary Syndrome: Thrombin Receptor Antagonist for Clinical Event

**Reduction in Acute Coronary Syndrome (TRACER)** 

Sponsor: Schering-Plough.

Total Cost: \$123,809 (target 13patients).

Role: Principal Investigator at the North Chicago VA Medical Center.

2008-2010 Project: A Multicenter, Double-Blind, Randomized Study to Establish

the Clinical Benefit and Safety of Vytorin (Ezetimibe/Simvastatin

Tablet) vs Simvastatin Monotherapy in High- Risk Subjects

Presenting With Acute Coronary Syndrome (IMProved Reduction of Outcomes: Vytorin Efficacy International Trial – IMPROVE IT)

Sponsor: Schering-Plough.

Total Cost: \$85.000 (target 17 patients).

Role: Principal Investigator at the North Chicago VA Medical Center.

2008-2010 Project: A Phase 3 Randomized, Double-blind, Parallel-group, Multi-

center Study of the Safety and Efficacy of Apixaban for Prophylaxis of Venous Thromboembolism in Acutely Ill Medical Subjects During and

Following Hospitalization Sponsor: Bristol Myers Squibb.

Total Cost: \$40,766.90 (target 5 patients).

Role: Principal Investigator at the North Chicago VA Medical Center.

2008-2010 Project: Stabilization Of pLaques usIng Darapladib" - A Clinical

Outcomes Study of Darapladib versus Placebo in Subjects Following Acute Coronary Syndrome to Compare the Incidence of Major Adverse Cardiovascular Events (MACE) (SOLID TIMI 52)

Sponsor: GlaxoSmithKline LLC. Total Cost: \$91,125. (target 10 patients).

Role: Principal Investigator at the North Chicago VA Medical Center.

**GIFTS** 

2004-2005 Project: Unrestricted Gift in Support of Industrial Partnership

Donor: Aventis Pharma Deutschland GmbH

Total Gift: \$50,000.

Role: Principal Investigator.

2004-2005 Project: Gift for Research and Education of First Responders to

**Victims of a Heart Attack** 

Donor: The Curtis I. Kossman Foundation

Total Cost: \$50,000.

Role: Principal Investigator.

2005-2006 Project: Gift for Research Activities

Donor: Pediatric Critical Care Specialists, P.C.

Total Cost: \$7,000.

Role: Principal Investigator.

2005-2007 Project: Gift for Research and Education of First Responders to

**Victims of a Heart Attack** 

Donor: The Curtis I. Kossman Foundation

Annual Gift: \$55,000. Role: Principal Investigator.

2008-2009 Project: Unrestricted Gift to Support Resuscitation Research

Donor: Michael G. Klug and Marie Kube

Total Gift: \$5,000.

Role: Principal Investigator.

2011 Project: Unrestricted Funds to Support Educational and Research

Activities

**Donor: ZOLL Medical Corporation** 

Total Gift: \$ 12,000

Role: Principal Investigator.

2013 Project: Unrestricted Funds for Cardiac Research

Donor: Monica Ply Total Gift: \$ 32,500

Role: Principal Investigator.